

IN THE TITLE:

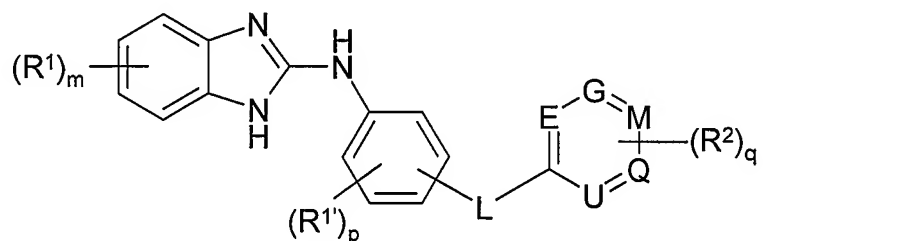
Please amend the title as follows,

~~Benzimidazolyl derivatives~~ Benzimidazoles

IN THE CLAIMS:

Please amend the claims as follows,

1. (Currently amended) A compound comprising formula I



wherein

R^1 , $R^{1'}$, R^2 each, independently of one another, are selected from the group consisting of Hal, A, OH, OA, SA, SO_2H , SO_2A , SO_3H , SO_3A , CN, NO_2 , NH_2 , NHA, NAA', NHCOA, CHO, $C(=O)A$, COOH, COOA, $CONH_2$, CONHA and CONAA',

L is selected from the group consisting of CH_2 , O and S,

E, G, M,

Q and U each, independently of one another, are selected from the group consisting of a C atom and an N atom, with the proviso that at least one and no more than one of E, G, M, Q or U is an N atom

A, A', independently of one another, are selected from the group consisting of unsubstituted or substituted alkyl having 1-10 C

atoms, unsubstituted or substituted cycloalkyl having 3-10 C atoms, unsubstituted or substituted alkoxyalkyl having 2-12 C atoms, unsubstituted or substituted aryl having 6-14 C atoms, unsubstituted or substituted arylalkyl having 7-15 C atoms, Hal is selected from the group consisting of F, Cl, Br and I, and m, p, q each, independently of one another, are 0, 1, 2, 3 or 4,

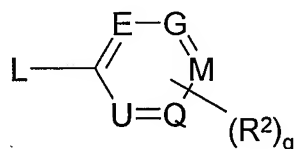
or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios.

2. (Currently amended) The compound according to Claim 1, wherein R^1 , independently of one another, is selected from the group consisting of A, Hal, CN, COOH, COOA, SO_2A , $C(=O)A$, NH_2 , NHA and NO_2 , and m is 1, 2 or 3, or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios.
3. (Currently amended) The compound according to Claim 1 wherein R^1 , independently of one another, is selected from the group consisting of methyl, ethyl, CF_3 , OCF_3 , F, Cl, Br, CN, COOH, $COOCH_3$, $COOCH_2CH_3$, SO_2CH_3 , NH_2 , $NHCH_3$, $NHCH_2CH_3$, NO_2 , and thiophen-2-ylcarbonyl, and m is 1, 2 or 3, or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios.

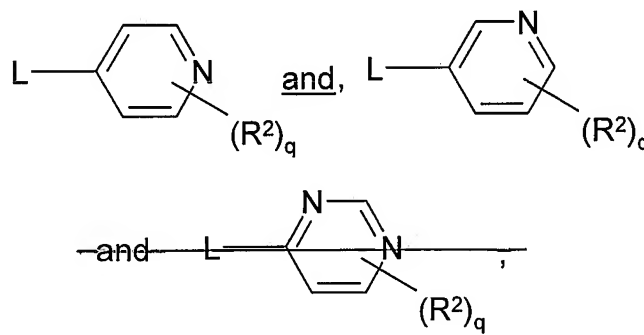
4. (Currently amended) The compound according to claim 1 wherein
R^{1'} is Hal or A,
p is 0 or 1,
or physiologically acceptable salts, solvates or stereoisomers thereof,
including mixtures thereof in all ratios.
5. (Cancelled)
6. (Currently amended) The compound according to claim 1 wherein
R² is selected from the group consisting of A, COOA, CONHA and
CONH₂, and
q is 0, 1 or 2,
or physiologically acceptable salts, solvates or stereoisomers thereof,
including mixtures thereof in all ratios.
7. (Currently amended) The compound according to claim 1 wherein
R¹, independently of one another, is selected from the group
consisting of Hal, alkyl, CN, COOH, COOalkyl, SO₂alkyl, NH₂,
NHalkyl, C(=O)alkyl, C(=O)heterocyclyl and NO₂,
m is 1, 2 or 3
R^{1'} is Hal or A
p is 0 or 1,
L is selected from the group consisting of O, S and ~~or~~ CH₂,
R² is selected from the group consisting of A, COOalkyl,
CONHalkyl and CONH₂, and
q is 0, 1 or 2,

or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios.

8. (Currently amended) The compound according to claim 1 wherein the group



in formula I is selected from the group consisting of



wherein L, R² and q have the meanings indicated in claim 1, or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios.

9. (Currently amended) The compound according to claim 1, selected from the group consisting of
(5-chloro-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;

[4-(pyridin-4-yloxy)phenyl](6-trifluoromethyl-1H-benzimidazol-2-yl)amine;
(6-methyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(5-chloro-4-methyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(4-bromo-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(4-bromo-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;
(5,6-dimethyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(5-chloro-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;
(5,6-dichloro-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(5,6-dichloro-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;
(5-chloro-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(5-chloro-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;
(4-methyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;
(4-chloro-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(4-chloro-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;
(4,5-dimethyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(5-chloro-6-methyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
(5-chloro-6-methyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;

(4,6-bis(trifluoromethyl)-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;

(4,6-bis(trifluoromethyl)-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;

[4-(pyridin-3-yloxy)phenyl](6-trifluoromethyl-1H-benzimidazol-2-yl)amine;

(6-methyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;

(4,5-dimethyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;

(5-chloro-4-methyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;

(4-methyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;

(5,6-dimethyl-1H-benzimidazol-2-yl)[4-(pyridin-3-yloxy)phenyl]amine;

~~(4-bromo-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(2,6-dimethylpyrimidin-4-yloxy)phenyl]amine;~~

N-methyl-4-[4-(bromotrifluoromethyl-1H-benzimidazol-2-ylamino)phenoxy]pyridine-2-carboxamide;

2-[4-(pyridin-4-yloxy)phenylamino]-3H-benzimidazole-5-carbonitrile;

~~[4-(2-amino-6-methylpyrimidin-4-yloxy)phenyl](4-bromo-6-trifluoromethyl-1H-benzimidazol-2-yl)amine;~~

~~(4-chloro-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(2,6-dimethylpyrimidin-4-yloxy)phenyl]amine;~~

~~[4-(2-amino-6-methylpyrimidin-4-yloxy)phenyl](4-chloro-6-trifluoromethyl-1H-benzimidazol-2-yl)amine;~~

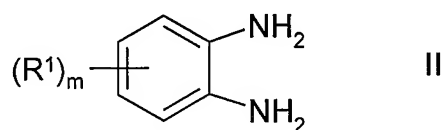
(6-nitro-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;

methyl 2-[4-(pyridin-4-yloxy)phenylamino]-3H-benzimidazole-5-carboxylate;

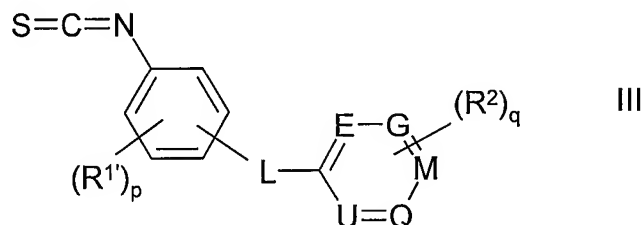
2-[4-(pyridin-4-yloxy)phenylamino]-3H-benzimidazole-5-carboxylic acid;
methyl
7-methanesulfonyl-2-[4-(pyridin-4-yloxy)phenylamino]-3H-benzimidazole-5-carboxylate;
(4-fluoro-6-trifluoromethyl-1H-benzimidazol-2-yl)[4-(pyridin-4-yloxy)phenyl]amine;
~~[4-(2,6-dimethylpyrimidin-4-yloxy)phenyl](4-fluoro-6-trifluoromethyl-1H-benzimidazol-2-yl)amine;~~
~~[4-(2-amino-6-methylpyrimidin-4-yloxy)phenyl](4-fluoro-6-trifluoromethyl-1H-benzimidazol-2-yl)amine;~~
N-methyl-4-{4-[6-(1-thiophen-2-ylmethanoyl)-1H-benzimidazol-2-yl-amino]phenoxy}pyridine-2-carboxamide; and
N²-[4-(pyridin-4-yloxy)phenyl]-3H-benzimidazole-2,5-diamine;
or physiologically acceptable salts, solvates or and stereoisomers thereof, including mixtures thereof in all ratios.

10. (Currently amended) A process for the preparation of a compound of the formula I according to claim 1 or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios, comprising reacting

a compound of the formula II



wherein R^1 and m have the meanings indicated in Claim 1,
with a compound of the formula III



wherein R^1 , L, E, G, M, Q, U, R^2 and q have the meanings indicated in Claim 1,

and optionally converting the compound of formula I into a salt.

11. (Currently amended) A pharmaceutical composition comprising at least one compound according to claim 1 or physiologically acceptable salts, solvates or stereoisomers thereof, including mixtures thereof in all ratios or optionally excipients or adjuvants.
12. (Previously presented) A method of treatment of diseases comprising inhibiting, regulating or modulating kinase signal transduction comprising administering to a patient in need thereof, a pharmaceutical composition according to claim 11.
13. (Currently amended) The method according to Claim 12, wherein said kinases are tyrosine kinases.
14. (Previously presented) The method according to Claim 13, wherein said tyrosine kinases are TIE-2.
- 15.-16. (Cancelled)

17. (Previously presented) The method according to Claim 12 wherein said disease comprises a solid tumour.
18. (Previously presented) The method according to Claim 17, wherein said solid tumour originates from the group consisting of brain tumour, tumour of the urogenital tract, tumour of the lymphatic system, stomach tumour, laryngeal tumour and lung tumour.
19. (Previously presented) The method according to Claim 17, wherein said solid tumour originates from the group consisting of monocytic leukaemia, lung adenocarcinoma, small cell lung carcinomas, pancreatic cancer, glioblastomas and breast carcinoma.
20. (Previously presented) The method according to Claim 12 wherein angiogenesis is implicated in said disease.
21. (Previously presented) The method according to Claim 20, wherein said disease is an ocular disease.
22. (Previously presented) The method according to Claim 12 wherein said disease is selected from the group consisting of retinal vascularisation, diabetic retinopathy, age-induced macular degeneration and inflammatory diseases.
23. (Previously presented) The method according to Claim 22, wherein said inflammatory disease originates from the group consisting of rheumatoid arthritis, psoriasis, contact dermatitis and delayed hypersensitivity reaction.

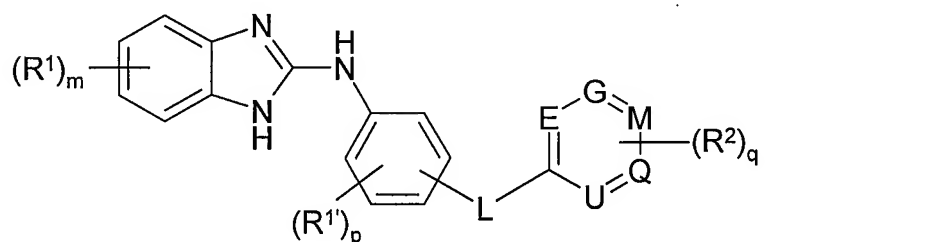
24. (Previously presented) The method according to Claim 12 wherein said disease involves bone pathologies, wherein said bone pathology originates from the group consisting of osteosarcoma, osteoarthritis and rickets.
25. (Previously presented) The pharmaceutical composition according to claim 11 comprising at least one additional active ingredient.
26. (Currently amended) A kit comprising separate packs of
 - (a) an effective amount of a compound according to Claim 1 or pharmaceutically usable derivatives, ~~selvates~~ and stereoisomers thereof, including mixtures thereof in all ratios, and
 - (b) an effective amount of ~~ad~~ an additional active ingredient.
27. (Previously presented) The method according to claim 12 wherein said pharmaceutical composition is administered in combination with a compound from the group consisting of 1) oestrogen receptor modulator, 2) androgen receptor modulator, 3) retinoid receptor modulator, 4) cytotoxic agent, 5) antiproliferative agent, 6) a prenyl-protein transferase inhibitor, 7) HMG-CoA reductase inhibitor, 8) HIV protease inhibitor, 9) reverse transcriptase inhibitor and 10) another angiogenesis inhibitor.
28. (Previously presented) The method according to claim 12 wherein said pharmaceutical composition is administered in combination with radiotherapy and a compound from the group consisting of 1) oestrogen receptor modulator, 2) androgen receptor modulator, 3) retinoid receptor

modulator, 4) cytotoxic agent, 5) antiproliferative agent, 6) prenyl-protein transferase inhibitor, 7) HMG-CoA reductase inhibitor, 8) HIV protease inhibitor, 9) reverse transcriptase inhibitor and 10) another angiogenesis inhibitor.

29. (Previously presented) The method according to claim 12 wherein said pharmaceutical composition is administered in combination with a growth-factor receptor inhibitor.
- 30.-31. (Cancelled)
32. (Previously presented) The method according to Claim 12 wherein said diseases are selected from the group consisting of hyperproliferative and non-hyperproliferative diseases.
33. (Previously presented) The method according to claim 12 wherein said disease is cancerous.
34. (Previously presented) The method according to claim 12 wherein said disease is non-cancerous.
35. (Previously presented) The method according to claim 34 wherein said non-cancerous diseases are selected from the group consisting of psoriasis, arthritis, inflammation, endometriosis, scarring, benign prostatic hyperplasia, immunological diseases, autoimmune diseases and immunodeficiency diseases.
36. (Previously presented) The method according to claim 33 wherein said cancerous diseases are selected from the group consisting of brain cancer, lung cancer, squamous cell cancer, bladder cancer, gastric cancer,

pancreatic cancer, hepatic cancer, renal cancer, colorectal cancer, breast cancer, head cancer, neck cancer, oesophageal cancer, gynaecological cancer, thyroid cancer, lymphoma, chronic leukaemia and acute leukaemia.

37. (Previously presented) A compound comprising formula I



wherein

R^1 , $R^{1'}$ each, independently of one another, are selected from the group consisting of Hal, A, OH, OA, SA, SO₂H, SO₂A, SO₃H, SO₃A, CN, NO₂, NH₂, NHA, NAA', NHCOA, CHO, C(=O)A, COOH, COOA, CONH₂, CONHA and CONAA', wherein A, A' independently of one another, are selected from the group consisting of unsubstituted or substituted alkyl having 1-10 C atoms, unsubstituted or substituted cycloalkyl having 3-10 C atoms, unsubstituted or substituted alkoxyalkyl having 2-12 C atoms, unsubstituted or substituted aryl having 6-14 C atoms, unsubstituted or substituted arylalkyl having 7-15 C atoms, R^2 is selected from the group consisting of Hal, A, OH, OA, SA, SO₂H, SO₂A, SO₃H, SO₃A, CN, NO₂, NH₂, NHA, NAA', NHCOA, CHO, C(=O)A, COOH, COOA, CONH₂, CONHA and CONAA', wherein A, A' independently of one another, are

selected from the group consisting of unsubstituted or substituted alkyl having 1-10 C atoms, unsubstituted or substituted cycloalkyl having 3-10 C atoms, unsubstituted or substituted alkoxyalkyl having 2-12 C atoms, unsubstituted or substituted aryl having 6-14 C atoms, unsubstituted or substituted arylalkyl having 7-15 C atoms, unsubstituted or substituted, saturated, unsaturated or aromatic heterocyclyl having 2-7 C atoms and 1-3 hetero atoms selected from the group consisting of N, O and S, or unsubstituted or substituted, saturated, unsaturated or aromatic heterocyclylalkyl having 3-10 C atoms and 1-3 hetero atoms selected from consisting of N, O and S,

L is selected from the group consisting of CH₂, O, and S,

E, G, M,

Q and U each, independently of one another, are selected from the group consisting of a C atom and ~~an~~ a N atom, with the proviso that at least one and no more than one of E, G, M, Q or U is an N atom,

Hal is selected from the group consisting of F, Cl, Br and I, and

m, p, q each, independently of one another, are 0, 1, 2, 3 or 4,

or physiologically acceptable salts, ~~solvates~~ or stereoisomers thereof, including mixtures thereof in all ratios.